[1032] Treatment continues in both arms in the absence of disease progression or unacceptable toxicity. Quality of life is assessed at baseline and day 1 of courses 2, 3, 4, 5, 6, 8, 10, and 12. Patients are followed at 1, 3, and 6 months and then every 6 months thereafter.

Example 33: Treatment for Bladder Cancer

[1033] Objective: Determine the acute toxicity of paclitaxel mid radiotherapy with or without a dolastatin derivative described herein in patients who have undergone prior transurethral bladder resection for muscle-invasive transitional cell carcinoma of the bladder.

[1034] Disease Characteristics: Histologically or cytologically is confirmed primary transitional cell carcinoma (TCC) of the bladder; histologic evidence of muscularis propria invasion; meets 1 of the following stage criteria: stage T2-4a; NX, N0, or N1; and M0 disease or clinical stage T1, grade 3/3 disease AND requires definitive local therapy; tumor involvement of the prostatic urethra allowed provided the following criteria are met: tumor is visibly completely resected; no evidence of stromal invasion of the prostate, no evidence of distant metastases by chest x-ray or CT scan AND abdominal/pelvic CT scan; has undergone transurethral bladder resection (as thorough as is judged safely possible) within the past 3-8 weeks, including bimanual examination with tumor mapping; sufficient tumor tissue available for HER2/neu analysis; not a candidate for radical cystectomy.

[1035] Study Design: This study is a non-randomized, multicenter study. Patients are assigned to 1 of 2 treatment groups according to HER2/neu status (HER2/neu 2+ or 3+ staining [group 1] vs HER2/neu 0 or 1+ staining [group 2]).

[1036] Group 1: Patients receive paclitaxel IV over 1 hour on days 1, 8, 15, 22, 29, 36, and 43 and a dolastatin derivative described herein via IV over 90 minutes on day 1 and then over 30 minutes on days 8, 15, 22, 29, 36, and 43. Patients also undergo radiotherapy once daily on days 1-5, 8-12, 15-19, 22-26, 29-33, 36-40, 43-47, and 50. Treatment continues in the absence of disease progression or unacceptable toxicity.

[1037] Group 2: Patients receive paclitaxel and undergo radiotherapy as in group 1. After completion of study treatment, patients are followed at 4-5 weeks, every 3 months for 1 year, every 4 months for 1 year, every 6 months for 3 years, and then annually thereafter.

Example 34: Treatment for Ovarian Cancer

[1038] Human Clinical Trial of the Safety and Efficacy of a Dolastatin Derivative described herein for Ovarian Cancer Therapy

[1039] Objective: Evaluate the safety and efficacy of a four week once weekly IV dosage of composition comprising a dolastatin derivative described herein in women with HER2-overexpressing ovarian cancer.

[1040] Study Design: This study is a non-randomized, open-label, 11 week, multicenter study. This study will evaluate the safety profile of four once weekly IV dosage, the MTD, PK and immunogenicity of trastuzumab-linked dolastatin derivative. Patients are assigned to a single group. Patients receive one dose of trastuzumab-linked dolastatin derivative once a week for 4 weeks. Trastuzumab-linked dolastatin derivative will be administered by IV infusion on Study Days 1, 8, 15, and 22. Urine samples will be taken on days 1 and 22.

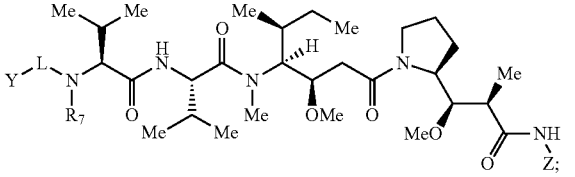
[1041] Blood Sampling Serial blood is drawn by direct vein puncture before and after administration of the dolastatin derivative. Venous blood samples (5 mL) for determination of serum concentrations are obtained at about 10 minutes prior to dosing and at approximately the following times after dosing: days 1, 2, 4, 5, 8, 15, 22, 36, 43 and 50. Each serum sample is divided into two aliquots. All serum samples are stored at −20° C. Serum samples are shipped on dry ice.

[1042] Treatment continues in the absence of disease progression or unacceptable toxicity. Quality of life is assessed at baseline and day 1 of courses 2, 3, 4, 5, 6, 8, 10, and 12. Patients me followed on days 29. 36, 43, and 50. Patients will be asked about adverse events. Patients will have an imaging scan and ECG to evaluate tumor size and heart function (day 43). At the termination of the study patients will have a physical exam day 50). Patients with evidence of disease regression may receive continued therapy until evidence of progression of disease is documented.

What is claimed is:

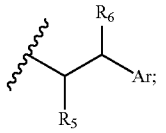
1. A compound, or salt thereof, comprising Formula (I):

(I)



wherein:

Z has the structure of:



R_5 is H, COR_8, C_1-C_6alkyl, or thiazole;

R_8 is OH or —NH-(alkylene-O)$_n$—NH_2;

R_6 is OH or H;

Ar is phenyl or pyridine;

R_7 is C_1-C_6alkyl or hydrogen;

Y and V are each selected from the group consisting of an hydroxylamine, methyl, aldehyde, protected aldehyde, ketone, protected ketone, thioester, ester, dicarbonyl, hydrazine, amidine, imine, diamine, azide, keto-amine, keto-alkyne, alkyne, cycloalkyne, and ene-dione;